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(FILE 'HOME' ENTERED AT 12:36:59 ON 22 OCT 2002)

FILE 'MEDLINE, CANCERLIT, CAPLUS, BIOTECHDS, EMBASE' ENTERED AT 12:37:19 ON 22 OCT 2002 1350013 S ADENOVIR? OR RETROVIR? OR VIRAL OR VIRUS L13884 S CATIONIC LIPID OR CATIONIC LIPOSOME L2 1043 S L2 AND L1 L3 210301 S DIAMETER L4L518 S L4 AND L3 11 DUP REM L5 (7 DUPLICATES REMOVED) L6 L7 699744 S BOUND OR ENCAPSULATED 66 S L7 AND L3 rs29 DUP REM L8 (37 DUPLICATES REMOVED) L9 4028 S CONDENS? AND L1 L10 57 S L10 AND L2 L1128 DUP REM L11 (29 DUPLICATES REMOVED) L12

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ANSWER 7 OF 11 BIOTECHDS COPYRIGHT 2002 THOMSON DERWENT AND ISI L6 1997-13269 BIOTECHDS ΑN Novel supramolecular assemblies for gene transfer; TIlipid-entrapped polycation-condensed DNA composition for use in lipofection and gene therapy (conference abstract)

ΑU Huang L

Univ.Pittsburgh CS

University of Pittsburgh, Department of Pharmacology, W1351 Biomedical LO Science Tower, Pittsburgh, PA 15261, USA.

Abstr.Pap.Am.Chem.Soc.; (1997) 213 Meet., Pt.2, PMSE306 SO ISSN: 0065-7727 CODEN: ACSRAL American Chemical Society, 213th ACS National Meeting, San Francisco, CA, 13-17 April, 1997.

DΤ Journal English LΑ

AΒ

The relatively non-toxic and efficient cationic liposome formulation DC-Chol-DOPE has been used in 2 separate clinical trials for immunotherapy of cancer and gene therapy of cystic fibrosis. Two types of novel condensed structure containing DNA polycation and lipids have been developed. These lipid-entrapped polycation-condensed DNA (LPD) particles are small (under 100 nm in diameter), monodispersed and colloidally stable. Transfection activity of LPD is similar to that of adeno virus vectors, and is 10- to 100-fold higher than that of first-generation cationic liposomes. LPD-I particles are cationic and used primarily in local and regional delivery routes. LPD-II particles are anionic and may be made target-specific by attaching specific ligands on the surface. Parenteral use of these novel particles for systemic gene transfer is under development. Recently, reconstituted chylomicron remnants have been used to solubilize DNA-cationic lipid complexes. This new non-virus vector induces high-level transgene expression in the liver. These formulations were discussed in terms of their efficiency, toxicity and uses in gene therapy. (0 ref)

L9 ANSWER 28 OF 29 CAPLUS COPYRIGHT 2002 ACS
AN 1994:571553 CAPLUS
DN 121:171553
TI Transfection of plant protoplasts with tobacc

TI Transfection of plant protoplasts with tobacco mosaic virus RNA by using electroporation, PEG and cationic liposome -mediated methods

AU Yuan, Tianhua; Wu, Jianhua; Gong, Zuxun; Yang, Jingping; Lin, Qishui CS Shanghai Inst. Biochem., Acad. Sin., Shanghai, 200031, Peop. Rep. China

SO Shengwu Huaxue Yu Shengwu Wuli Xuebao (1994), 26(1), 7-13 CODEN: SHWPAU; ISSN: 0582-9879

DT Journal

Chinese LΑ Tobacco mosaic virus RNA was introduced into protoplasts from AΒ tobacco (Nicotiana tabacum L. cv. Bright Yellow) and Chinese cabbage (Brassica chinensis) by electroporation, PEG treatments and cationic liposome-mediated methods. The results indicated that although both of electroporation and PEG treatment methods could introduce TMV-RNA into protoplasts, cationic liposome-mediated methods could enhance infection significantly. The min. amt. of TMV-RNA necessary for transfection of protoplasts decreased down to 10 times, when the TMV-RNA mols. were encapsulated before introduction into protoplasts by electroporation or PEG treatment. The results also showed that TMV could multiply inside protoplasts and reach the max. after 48 h of introduction of TMV-RNA. SDS-PAGE of the roughly extd. solns. of transfected protoplasts at 48 h after introduction of TMV-RNA showed that a 17.5 kd band, the mol. wt. of which is equal to TMV coat protein clearly appeared and besides, there also a 50-55 kd protein band could be obsd. from these transfected protoplasts.

ANSWER 26 OF 29 BIOTECHDS COPYRIGHT 2002 THOMSON DERWENT AND ISI L9 1995-12639 BIOTECHDS AN Adeno viral mediated cell transfection; ΤI receptor-mediated gene transfer to e.g. human cell in vivo using wild-type or inactivated virus or empty capsid, optionally with lipofection, e.g. for gene therapy Seth P; Crystal R G; Rosenfeld M; Yoshimura K; Jessee J A ΑU U.S.Dep.Health-Hum.Serv.; Life-Technol. PAWO 9521259 10 Aug 1995 PΙ .WO 1995-US924 24 Jan 1995 ΑI US 1994-191669 4 Feb 1994 PRAI Patent DTEnglish LΑ OS WPI: 1995-283779 [37] A new method for introducing a nucleic acid (NA) (DNA, RNA or peptide AB nucleic acid) into a eukaryotic cell (mammal, bird or fish, especially ungulate, cat, dog or human) in vivo or in vitro involves contact of the cell with the NA and an adeno virus (Ad), where the NA is not bound to a molecule. The cell is contacted with the Ad less than 8 hr (especially less than 2 hr) before or after contact with NA. adeno virus (wild-type or modified to become replication-deficient) is present at 20-2,000 pfu/cell. The modification is an insertion, rearrangement, deletion, replacement, methylation, demethylation or mutagenesis, and alters cell binding, endosomal lysis or intracellular targeting. The Ad may be in empty capsid form, or inactivated. A cationic agent, e.g. a polycarbene, 1,5-dimethyl-1,5diazaundecamethylene polymethobromide or a cationic liposome may be mixed with the NA and used to effect entry into the cell. The Ad transfers cargo molecules into the cell nucleus by receptor-mediated endocytosis. Unlike previous methods, this method may be used in vivo, and overcomes fusion difficulties, nucleic acid degradation and toxicity problems. (66pp)

L9 ANSWER 25 OF 29 MEDLINE DUPLICATE 15

AN 97261594 MEDLINE

DN 97261594 PubMed ID: 9107521

TI Lipidic vector systems for gene transfer.

AU Lee R J; Huang L

CS Endocyte, Inc., West Lafayette, IN 47906, USA.. leer@endocyte.wintek.com

SO CRITICAL REVIEWS IN THERAPEUTIC DRUG CARRIER SYSTEMS, (1997) 14 (2) 173-206. Ref: 67
Journal code: 8511159. ISSN: 0743-4863.

CY United States

DT Journal; Article; (JOURNAL ARTICLE)
General Review; (REVIEW)
(REVIEW, TUTORIAL)

LA English

FS Priority Journals

EM 199707

ED Entered STN: 19970716 Last Updated on STN: 19970716 Entered Medline: 19970702

Clinical application of gene therapy depends on the development of AΒ suitable gene transfer vehicles (vectors). Although generally not as efficient as viral vectors, nonviral systems such as lipidic vectors have the potential advantages of being less toxic, nonrestrictive in cargo DNA size, potentially targetable, and easy to produce in relatively large amounts. More important, lipidic vectors generally lack immunogenicity, allowing repeated in vivo transfection using the same vector. In this paper, we will attempt to summarize some of the recent advances in lipidic gene delivery vectors. Three types of lipidic gene transfer vectors are described: 1) DNA/cationic liposome complexes, 2) DNA encapsulated in neutral or anionic liposomes, and 3) liposome-entrapped, polycation-condensed DNA (LPDI and LPDII). We review the various factors affecting vector structure and gene delivery efficiency, and we discuss the possible mechanisms of gene transfer and their implications in vector design.

DUPLICATE 7 MEDLINE ANSWER 15 OF 29 L9 1999291932 MEDLINE AN99291932 PubMed ID: 10365810 DN Gene transfer to the rat biliary tract with the HVJ-cationic TIliposome method. Uehara T; Honda K; Hatano E; Terao R; Iimuro Y; Yamamoto N; Yamamoto M; ΑU Kaneda Y; Yamaoka Y Department of Gastroenterological Surgery, Kyoto University, Graduate CS School of Medicine, Japan.. tetsuzo@kuhp.kyoto-u.ac.jp JOURNAL OF HEPATOLOGY, (1999 May) 30 (5) 836-42. Journal code: 8503886. ISSN: 0168-8278. SO CY Denmark Journal; Article; (JOURNAL ARTICLE) DTEnglish LΑ Priority Journals FS EM 199907 Entered STN: 19990806 ED Last Updated on STN: 19990806 Entered Medline: 19990723 BACKGROUND/AIMS: The ability to transfer foreign genes into the biliary AB tract would be useful for the treatment of biliary tract diseases, including cancer, cystic fibrosis and other genetic diseases. To introduce a foreign gene precisely into the rat biliary epithelial cells, we developed a new technique, inserting a polyethylene catheter into the common bile duct through the papilla of Vater by use of a fusigenic cationic liposome with hemagglutinating virus of Japan (HVJ-cationic liposome). METHODS: Transfection efficiency was estimated with the use of FITColigonucleotides (FITC-ODNs) and cDNA of beta-galactosidase (pCAG-lacZ). RESULTS: FITC-ODNs encapsulated in HVJ-cationic liposome were effectively transfected into cell nuclei of human cholangiocellular carcinoma in vitro after a 30-min incubation as compared with the simple application of naked FITC-ODNs. After in vivo injection of FITC-ODNs using the HVJ-cationic liposome method through the papilla of Vater, fluorescence accumulation was observed only in the epithelial cells of the biliary tract, but not in the parenchymal cells of the liver. Beta-galactosidase expression was observed in the biliary epithelial cells 3 days after the transfection of pCAG-lacZ and

was also detected at 14 days, but not at 28 days, without obvious

-mediated gene transfer to the biliary tract via the papilla of Vater is a minimally-invasive and an effective gene-delivery method for site-specific targeting to the epithelial cells of the biliary tract, which could be

cytotoxicity. CONCLUSIONS: HVJ-cationic liposome

applied to the treatment of human biliary tract diseases.

ANSWER 8 OF 29 CAPLUS COPYRIGHT 2002 ACS L9 2000:145054 CAPLUS ANDN 132:176589 Cationic complexes of polymer-modified adenovirus vectors ΤI Wadsworth, Samuel C.; O'Riordan, Catherine E. IN Genzyme Corporation, USA PA PCT Int. Appl., 73 pp. SO CODEN: PIXXD2 DT Patent LΑ English FAN.CNT 1 APPLICATION NO. DATE PATENT NO. KIND DATE \_\_\_\_\_ \_\_\_ \_\_\_\_\_ WO 1999-US19162 19990823 20000302 WO 2000011202 A1 PΙ W: AU, CA, JP, US RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE AU 1999-56857 19990823 A1 20000314 AU 9956857 20010620 EP 1999-943838 19990823 EP 1108048 Α1 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI JP 2000-566454 19990823 20020730 Т2 JP 2002523054 19980824 PRAI US 1998-97653P Ρ 19990823 WO 1999-US19162 W An adenovirus complex including a complex of a cationic mol. and AΒ of an adenovirus having at least one polyalkylene glycol polymer bound thereto. The polyalkylene glycol polymer includes, but is not limited to, polyethylene glycol, methoxypolyethylene glycol, polymethylethylene glycol, polyhydroxypropylene glycol, polypropylene glycol, and polymethylpropylene glycol. Mol. wts. for the polymer range from 200 to 20,000 Daltons, with 2000 to 12,000 Daltons being preferred. The adenovirus is a preferably recombinant adenoviral vector such as a recombinant viral vector contg. a transgene. The polymer is bound, directly or indirectly, to the virus particle by covalent or noncovalent means. The cationic mol. is preferably a cationic polymer, such as DEAE-Dextran, or a cationic lipid. A compn. contg. the adenovirus complex and a carrier is also disclosed. The complexes of the invention exhibit reduced immunogenicity.

L9 ANSWER 1 OF 29 MEDLINE DUPLICATE 1

AN 2002253956 MEDLINE

DN 21969385 PubMed ID: 11973632

TI Characterisation of LMD virus-like nanoparticles self-assembled from cationic liposomes, adenovirus core peptide mu and plasmid

- AU Tagawa T; Manvell M; Brown N; Keller M; Perouzel E; Murray K D; Harbottle R P; Tecle M; Booy F; Brahimi-Horn M C; Coutelle C; Lemoine N R; Alton E W F W; Miller A D
- CS Imperial College Genetic Therapies Centre, Department of Chemistry, Imperial College of Science, Technology and Medicine, London, UK.
- SO GENE THERAPY, (2002 May) 9 (9) 564-76. Journal code: 9421525. ISSN: 0969-7128.
- CY England: United Kingdom
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals
- EM 200205
- ED Entered STN: 20020508
  Last Updated on STN: 20020514
  Entered Medline: 20020513
- Liposome:mu:DNA (LMD) is a ternary nucleic acid delivery system built AΒ around the mu peptide associated with the condensed core complex of the adenovirus. LMD is prepared by precondensing plasmid DNA (D) with mu peptide (M) in a 1:0.6 (w/w) ratio and then combining these mu:DNA (MD) complexes with extruded cationic liposomes (L) resulting in a final lipid:mu:DNA ratio of 12:0.6:1 (w/w/w). Correct buffer conditions, reagent concentrations and rates of mixing are all crucial to success. However, once optimal conditions are established, homogeneous LMD particles (120 +/- 30 nm) will result that each appear to comprise an MD particle encapsulated within a cationic bilammellar liposome. LMD particles can be formulated reproducibly, they are amenable to long-term storage (>1 month) at -80 degrees C and are stable to aggregation at a plasmid DNA concentration up to 5 mg/ml (15 mM nucleotide concentration). Furthermore, LMD transfections are significantly more time and dose efficient in vitro than cationic liposome-plasmid DNA (LD) transfections. Transfection times as short as 10 min and plasmid DNA doses as low as 0.001 microg/well result in significant gene expression. LMD transfections will also take place in the presence of biological fluids (eg up to 100% serum) giving 15-25% the level of gene expression observed in the absence of serum. Results from confocal microscopy experiments using fluorescent-labelled LMD particles suggest that endocytosis is not a significant barrier to LMD transfection, although the nuclear membrane still is. We also confirm that topical lung transfection in vivo by LMD is at least equal in absolute terms with transfection mediated by  $\label{eq:GL-67:DOPE:DMPE-PEG(5000)} $$(1:2:0.05\ m/m/m)$, an accepted 'gold-standard' $$$ non-viral vector system for topical lung transfection, and is in fact at least six-fold more dose efficient. All these features make LMD an important new non-viral vector platform system from which to derive tailor-made non-viral delivery systems by a process of systematic modular upgrading.

ANSWER 7 OF 28 BIOTECHDS COPYRIGHT 2002 THOMSON DERWENT AND ISI L12 2001-14354 BIOTECHDS ΑN Nucleic acid delivery complex for delivering nucleic acids to cells such TIas neuronal, cancer, epithelial cells, comprises cationic lipid/protein/nucleic acid complex comprising viral packaging proteins; nucleic acid vaccine delivery and gene therapy Tagawa T; Miller A D; Perouzel E; Murray K; Manvell M; Alton E; Matthews ΑU D; Russell W Mitsubishi-Tokyo-Pharmaceuticals PA LO Tokyo, Japan. WO 2001048233 5 Jul 2001 PΙ WO 2000-GB4767 12 Dec 2000 ΑI GB 1999-30533 23 Dec 1999 PRAI Patent DTEnglish LΑ WPI: 2001-441719 [47] OS A non-viral nucleic acid delivery vector (I) comprising a AΒ condensed polypeptide/nucleic acid complex and a cationic lipid, is claimed, where the complex comprises a nucleic acid sequence of interest and one or more virus nucleic acid packaging proteins, or their derivatives capable of binding to and condensing the nucleic acid of interest, which is heterologous to the protein. Also claimed are: a condensed protein/nucleic acid complex (II); producing (I) by contacting the nucleic acid of interest with a virus nucleic acid packaging protein or its derivative and contacting the nucleic acid/protein complex formed with a cationic lipid; and use of a virus nucleic acid packaging protein or its derivative in the manufacture of (I). (I) and (II) are used to introduce a nucleic acid of interest into a eukaryotic cell, especially a neuronal, cancer or epithelium cell. and (II) can be used in gene therapy, nucleic acid vaccine delivery and in vitro transfection studies. (71pp)